PATENT

In re application of:	Curtis, Rory A.J.		
Application No.:	10/074,547	Group No.:	1647
Filed:	February 12, 2002	Examiner:	Kapust, Rachel B.
For:	25466, A HUMAN TRANSPORT	ER FAMILY M	MEMBER AND USES THEREFOR

Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

DECLARATION UNDER 37 C.F.R. § 1.131

Sir:

I, Rory A.J. Curtis, hereby declare and state:

- 1. I am the inventor of the subject matter described and claimed in the above-identified application.
- 2. I was in possession of the complete nucleic acid sequence of 25466 in the United States and had determined its identity as a monocarboxylate transporter before January 30, 2001.
- 3. Evidence is provided by the following:
 - a) Prior to January 30, 2001, I had completed the sequencing of full length human 25466. Exhibit A is a copy of an electronic printout of the nucleic acid sequence from Millennium's sequence database. The printout bears the automatically embedded date on which I entered the sequence into the database. On the original printout, this sequence bears a date prior to January 30, 2001. In accordance with accepted practice, the dates on the copy of the electronic printout have been masked (M.P.E.P. § 715.07).

The top sections of the printout bear reference information for accessing the 25466

CERTIFICATION UNDER 37 C.F.R. SECTIONS 1.8(a) and 1.10* I hereby certify that, on the date shown below, this correspondence is being: MAILING deposited with the United States Postal Service in an envelope addressed to the Commissioner for Patents, P.O. Box \square 1450, Alexandria, VA 22313-1450. 37 C.F.R. SECTION 1.10* 37 C.F.R. SECTION 1.8(a) as "Express Mail Post Office to Addressee" with sufficient postage as first class mail. \square Mailing Label No. TRANSMISSION transmitted by facsimile to the Patent and Trademark Office. Sean Hunziker/Beverly Sotiropoulos July 8, 2004 Date:

*WARNING:Each paper or fee filed by "Express Mail" must have the number of the "Express Mail" mailing label placed thereon prior to mailing. 37 C.F.R. section 1.10(b). "Since the filing of correspondence under section 1.10 without the Express Mail mailing label thereon is an oversight that can be avoided by the exercise of reasonable care, requests for waiver of this requirement will not be granted on petition." Notice of Oct. 24, 1996, 60 Fed. Reg. 56,439, at 56,442.

(type or print name of person certifying)

(Page 1 of 2)

Practitioner's Docket No. MPI01-019P1RNM

sequence in the Millennium database. The remainder of the printout bears the entire 25466 nucleotide sequence ("Fbh25466pat"), known in the application as SEQ ID NO:1.

Prior to January 30, 2001, I had determined that the 25466 nucleotide sequence of SEQ ID NO:1 encodes a monocarboxylate transporter based on the results of analyses of the nucleotide sequence and the encoded polypeptide. Exhibits B, C and D are evidence of this determination. Exhibit B is a copy of the first page of a BLAST analysis of SEQ ID NO:1. The top hit in this BLAST result is human monocarboxylate transporter 3 (Genbank Accession No. U81800). Exhibit C is a copy of several analyses of the 25466 polypeptide, with the sequence displayed on page 1. Page 4 of Exhibit C shows the alignment of a portion of the polypeptide with the MCT (monocarboxylate transporter) domain in Pfam. Exhibit D is a pairwise alignment of the 25466 polypeptide with human monocarboxylate transporter 4 (SwissProt Accession No. O15374). I performed my analyses and reached the conclusion that SEQ ID NO:1 encodes a monocarboxylate transporter prior to January 30, 2001.

The original printouts of Exhibits B and C bear automatically embedded dates on which I performed these analyses. In accordance with accepted practice, the dates on the copies of the electronic printouts have been masked (M.P.E.P. § 715.07).

I hereby declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code, and that such willful false statements may jeopardize the validity of the application or any patent issued thereon.

Rory A.J. Curtis

Date

Sample Information

name: Fbh25466pat

type: Plain DNA Sequence

submitted on:
NC project: Bayer
species: Human

tissue: unspecified internal id: 107698647 restrictions: none

Sequences Information

creation method: Loading

name: Load created on:

created by: Rory Curtis
accession number: 107698648

> Fbh25466pat - Import - complete

1	GTCGACCCAC	GCGTCCGCAA	GAGTGTGCAT	GTGAGGTGAC	TGCATTTTTT	TTCCCTGCCA
61	AACCAGAATT	AGCCGGTATA	GGAATGAACG	AGCATGAAGA	TTTGAAATTG	CTCCGATTGG
121	AAGGAAGCCC	AGGTTAGGTT	TGGGCACCTC	CAAACGCACC	CGTTTTAAAG	CCACCTGGAC
181	TGAGGCGTCG	AGCTTTCAGC	TCCACCAAAC	GCTCACCTGG	CCTGGCAGCG	AGCGGCGGAA
241	GAGCCCGGGA	GCCCTCACA	GAGCGCACCG	AGCCGGGCGG	AGAGCTGAGC	CGCAGGCACC
301	CGCGTCTCCA	GGATGATAGG	CGACATTGCA	ACAAATCTCT	ACACCCAGCA	GCTCAGGGGG
361	CTCCAAGCAG	AGCAGCAAGT	TCGAGGATCC	GGGCGTGGAG	CCGAGTGAGG	CCGCAGCCCA
421	GCGGGCCTCG	GGCGAAAAAT	CTTGGAAAAT	GTATACCAGT	CATGAAGATA	TTGGGTATGA
481	TTTTGAAGAT	GGCCCCAAAG	ACAAAAAGAC	ACTGAAGCCC	CACCCAAACA	TTGATGGCGG
541	ATGGGCTTGG	ATGATGGTGC	TCTCCTCTTT	CTTTGTGCAC	ATCCTCATCA	TGGGCTCCCA
601	GATGGCCCTG	GGTGTCCTCA	ACGTGGAATG	GCTGGAAGAA	TTCCACCAGA	GCCGCGGCCT
661	GACCGCCTGG	GTCAGCTCCC	TCAGCATGGG	CATCACCTTG	ATAGTGGGCC	CTTTCATCGG
721	CTTGTTCATT	AACACCTGTG	GGTGCCGCCA	GACTGCGATC	ATTGGAGGGC	TCGTCAACTC
781	CCTGGGCTGG	GTGTTGAGTG	CCTATGCTGC	AAACGTGCAT	TATCTCTTCA	TTACTTTTGG
841	AGTCGCAGCT	GGCCTGGGCA	GCGGGATGGC	CTACCTGCCA	${\tt GCGGTGGTCA}$	TGGTGGGCAG
901	GTATTTCCAG	AAGAGACGCG	CCCTCGCCCA	GGGCCTCAGC	ACCACGGGGA	CCGGATTCGG
961	TACGTTCCTA	ATGACTGTGC	TGCTGAAGTA	CCTGTGCGCA	GAGTACGGCT	GGAGGAATGC
1021	CATGTTGATC	CAAGGTGCCG	TTTCCCTAAA	CCTGTGTGTT	TGTGGGGCGC	TCATGAGGCC
1081	CCTCTCTCCT	GGTAAAAACC	CAAACGACCC	AGGAGAGAAA	GATGTGCGTG	GCCTGCCAGC
1141	GCACTCCACA	GAATCTGTGA	AGTCAACTGG	ACAGCAGGGA	AGAACAGAAG	AGAAGGATGG
1201	TGGGCTCGGG	AACGAGGAGA	CCCTCTGCGA	CCTGCAÁGCC	CAGGAGTGCC	CCGATCAGGC
1261	CGGGCACAGG	AAGAACATGT	GTGCCCTCCG	GATTCTGAAG	ACTGTCAGCT	GGCTCACCAT
1321	GAGAGTCAGG	AAGGGCTTCG	AGGACTGGTA	TTCGGGCTAC	TTTGGGACAG	CCTCTCTATT
1381 _.	TACAAATCGA	ATGTTTGTAG	CCTTTATTTT	CTGGGCTTTG	TTTGCATACA	GCAGCTTTGT
1441	CATCCCCTTC	ATTCACCTCC	CAGAAATCGT	CAATTTGTAT	AACTTATCGG	AGCAAAACGA
1501	CGTTTTCCCT	CTGACGTCAA	TTATAGCAAT	AGTTCACATC	TTTGGAAAAG	TGATCCTGGG

Exhibit A to Accompany Declaration under 37 CFR §1.131 for US Appln. No. 10/074,547

	1561	ССФСАТАССС	GACTTGCCTT	GCATTAGTGT	ТТССААТСТС	TTCCTGTTGG	CCAACTTCAC	
	1621			TTCTGCCGTT				
	1681						TGACTGAAGA	
	1741			TGGCCAATGC				
	1801	**					AAAAATATGA	
	1861						TACTTATTCA	
	1921						CACATGTTTA	
	1921			TAGGTTTCAT				
·				GACGTGGGAA				
	2041			GCCTCATTTG				
	2101			AGGACTACAT				
4	2161							
	2221			TAGAAATGCC AGCTTTTAAA				
	2281							
	2341						AGCTTACAGA	
	2401			GAATATGACT				
•	2461						ACAATTTAGT	
	2521						GGATTGTGAC	
	2581						GCTATCCAGT	
	2641			AAAACCTTAA				
	2701			CAACATTAAA				
	2761						TTTTTTCTGA	
	2821						AAAGTTAATA	
	2881						TTTGTGAGGA	
	2941						GAGTAGTGAT	
	3001						TGCCTGAGCG	
	3061						CTGTGATCCA	•
	3121				•		CAAACGTTTT	
	3181						ACTTGTTTAC	
	3241	***************************************					CACACAAATC	
	3301						AGCCCATTCT	
•	3361						AACAATGCCA	
	3421			AGCAGTTTCA				
	3481						TCCTGCTCTA	
	3541						GGCAGCTAAT	
	3601						AATGACTCGG	
	3661						TGGTTCAACA	
	3721						GCTCTCTATT	
	3781	TTGATAAAGA	TAATCTTAAA	GCTGAGGCAA	TGCTCCCTCC	CCTATCTCTC	TCTGTGTAAT	
	3841	TTACCATAGA	ATTAGGATGA	TTAGATTGAA	ACACATGTTG	TATGTTTTAA	AAACTACATT	
	3901			•			TAAATATTTT	•
	3961	AAATGGTGCT	ATAGCCAATG	TATTTGAATG	CTTGCACTGC	TGGTTGTGTA	TCATCAATAT	
	4021	GAACTTTTTA	TCCAATGACT	CAACTCTAAT	TACATCTAAG	TTAGACTTGC	TCACGTTCAG	
	4081	TTTGTACAGT	TGTGTGTTGA	CTTACTATGT	TTTGAAAGTG	GTGACTTCTA	CCGAATGAGT	
	4141	GGAAGTTCCC	ATTGTCAAAA	AAAATAAAGA	CCTGCTTGCA	GTATTCATGT	TGACAACAGA	
	4201	GTAAAAGAGA	ATACTGTAAA	GAATTACTGC	AAATATTTCC	TGTTTATGTT	ATTTGCCGTT	
	4261						TCGTTTTCCC	
	4321	CTCATGTATC	CAAGTAATTT	TTATTTACAT	ACAACTAAAT	AAATGTTGTC	CTCTTTGAAA	

Fbh25466pat(4419 bases) -Nucleic Acid Db Check - All Database Hits

gb|U81800|HSU81800 Homo sapiens monocarboxylate transport... 384 8.4e-20 1 gb|U15685|GGU15685 Gallus gallus retinal epithelial membr... 376 4.0e-19 1 gb AJ238706 DME238706 Drosophila melanogaster mRNA for ca... 3.9e-12 292 gb AF178954 AF178954 Mus musculus monocarboxylate transpo... 280 4.0e-11 gb|AL009193|DMC103B4 Drosophila melanogaster cosmid clone... 292 2.4e-10 gb|AE003423|AE003423 Drosophila melanogaster genomic scaf... 292 1.3e-09 2 gb AE003469 AE003469 Drosophila melanogaster genomic scaf... 271 3.5e-09 2 gb AR083262 AR083262 Sequence 9 from patent US 5976837. 231 4.9e-07 1 >gb|U81800|HSU81800 Homo sapiens monocarboxylate transporter (MCT3) mRNA, complete cds.

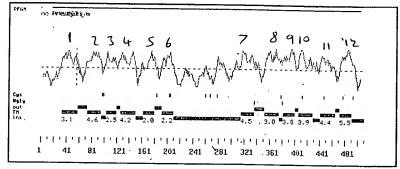
Length = 1982

Plus Strand HSPs:

		(108.6 bits), Expect = 8.4e-20, P = 8.4e-20 = 288/552 (52%), Positives = 288/552 (52%), Strand = Plus / Pl	.us
Query:	532	TGATGGCGGATGGCTTGGATGATGGTGCTCTCTTTTTTTT	591
Sbjct:	110	TGACGGCGGCTGGGCCGTGCTCTCGGCTGTTCGTCATCACTGGCTTCTCCTA	169
Query:	592	GGGCTCCCAGATGGCCCTGGGTGTCCTCAACGTGGAATGGCTGGAAGAATTCCACCAGAG	651
Sbjct:	170	CGCCTTCCCCAAGGCCGTCAGTGTCTTCTTCAAGGAGCTCATACAGGAGTTTGGGATCGG	229
Query:	652	CCGCGGCCTGACCGCCTGGGTCAGCTCCCTCAGCATGGGCATCACCTTGATAGTGGGCCC	711
Sbjct:	230	CTACAGCGACACGCTGGATCTCCTCCATCCTGCTGGCCATGCTCTACGGGACAGGTCC	289
Query:	712	TTTCATCGGCTTGTTCATTAACACCTGTGGGTGCCGCCAGACTGCGATCATTGGAGGGCT	771
Sbjct:	290	GCTCTGCAGTGTGCGTGAACCGCTTTGGCTGCCGGCCCGTCATGCTTGTGGGGGGTCT	349
Query:		CGTCAACTCCCTGGGCTGGTGTTGAGTGCCTATGCTGCAAACGTGCATTATCTCTTCAT	831
Sbjct:		CTTTGCGTCGCTGGGCATGGTGGCTGCGTCCTTTTGCCGGAGCATCATCCAGGTCTACCT	409
Query:	832	TACTTTTGGAGTCGCAGCTGGCCTGGGCAGCGGGATGGCCTACCTGCCAGCGGTGGTCAT	891
Sbjct:	410	CACCACTGGGGTCATCACGGGGTTGGGTTTGGCACTCAACTTCCAGCCCTCGCTCATCAT	469
Query:	892	GGTGGGCAGGTATTTCCAGAAGAGACGCCCCTCGCCCAGGGCCTCAGCACCACGGGGAC	951
Sbjct:	470	${\tt GCTGAACCGCTACTTCAGCAAGCGGCGCCCCATGGCCAACGGGCTGGCGGCAGCAGGTAG}$	529

Back to orfanal.cgi

Analysis of 25466 (510 aa)



Vo Signal Sequence
12 Transmembrane
Obnains
Cytoplasmic N- and
C-terminals

>25466
HYTSHEDIGYDPEDGPKDKKTLKPHPHIDGGWAWMMYLSSPFVHILIMGSQMALGYLNVE
WLEEPHOSRGLTAWVSSLSHGITLIVOPPIGLFINTCOCROTALIGGLUNSLGWULSAYA
MUNTHLFITGYAAGLGSGAWAYLPAVWMGSPYGKRALAQGISTTOTFOFTHWTULK
YLCASYGWRIAMALIQGAVSLNICVCGALHRPLSFCKINNDDGEKDVRGLPAHSTESYKST
YLCASYGWRIAMALIQGAVSLNICVCGALHRPLSFCKINNDDGEKDVRGLPAHSTESYKST
YLCASYGWRIAMALIQGAVSLNICVCGALHRPLSFCKINNDDGEKDVRGLPAHSTESYKST
YLCASYGWRIAMALIQGAVSLNICVCGALHRPLSFCKINNDDGEKDVRGLPAHSTESYKST
YSGYGTASLPTNRHWYAFIFMALFAYSSFYIPPIHLPEIVHLYMLSEGNDVFPLTSIIA
YHGIGKYLUGYADLPGISWNYVPLLAMFTULSFFILDHWTYAGLAYCALIGFSG
YFSLMPWYTEDLVGIEHLANAYGIIICANGISALLGPPPAGMIYDITQKYDPSFYICGLL
YHIGILFLIQPCIRIIEOSRRKYMDGAHY

PSORT Prediction of Protein Localization

Gavel: prediction of cleavage sites for mitochondrial preseq cleavage site motif not found

NUCDISC: discrimination of nuclear localization signals pat4: none pat7: PKDKRTL (5) at 16 bipartite: none content of basic residues: 6.5% NLS Score: -0.04

Final Results (k = 9/23):

66.7 %: endoplasmic reticulum 22.2 %: mitochondrial 11.1 %: nuclear

prediction for 29466 is end (k=9)

Start	End	Feature	Seq
387	408	Leucine zipper pattern (PS00029)	LANFILVLSILPLMHTYAGL

Signal Peptide Predictions for 25466

Method	Predict	Score	Mat@
SignalP (eukaryote)	MAYBE		56

N

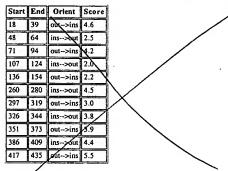
Note: amino-terminal 70aa used for signal peptide prediction

Transmembrane Segments Predicted by MEMSAT

Start	End	Orient	Score	
33	57	ins>out	3.1	TMI
73	94	out>ins	4.6	TMZ
103	119	ins>out	2.5	TM3
126	149	out>ins	4.2	TM4
162	179	ins>out	2.0	TIMS
191	209	out>ins	2.2	TMB
315	335	ins>out	4.5	TM7
352	374	out>ins	3.0	TIME
381	399	ins>out	3.8	TM9
406	428	out->ins	3.9	This
441	464	ins>out	4.4	TIM
472	490	out>ins	5.5	TMIL

>25466
MYTSHEDIGYDPEDGPRDKRTLKPHPHIDGGWAMGEN/LSSFFVHILIMGSGMALGVLNVE
WLBEFHGSRGLTANVSSLSKGTTLIVGPFIGLETTCGCRGTATIGGLNISLGMYLSAYA
ANNTHLFITGVÄRGLIGSGFAVLFAVDYNGSFYGRRALAGGESTTGTGFTFHTTHTVLLK
YLCARYGMRAMALICGAVSLNLCVCGALWRFLSPGKNENDGGERUVGGLPAHSTESVKST
YLCARYGMRAMALICGAVSLNLCVCGALWRFLSPGKNENDGARIKTVSMLTHRWKGPEDM
YSCYGTASLFYNNHFYAFIFALFAVSFVIPFIHLPEIVNLTYMLSEGNDVFFLTSIIA
TWHIEGKYLIGVIADLPGISVMYVELLARFTUVSSFFILPHTTAGLAVICALIGFSSG
YFSLEPYUTEDLWGIBHLAMAYGIIICANGISALGPPFAGMYDTTOKYDPSFYICGLL
YNIGILFLUGADLPGISVMYELLARFTUNSFFILPHTTAGLAVICALIGFSSG
YFSLEPYUTEDLWGIBHLAMAYGIIICANGISALGPPFAGMYDTTOKYDPSFYICGLL
YNIGILFLUGARGFAGMAY

Transmembrane segments for presumed mature peptide



>23466 ature

>ZIAGE ATURE

>Z

Prosite Pattern Matches for 25466

Prosite version: Release 12.2 of February 1995

>PS00001|PD0C00001|ASN_GLYCOSYLATION N-glycosylation site.

Query: 345 NLSE 348 Query: 389 NPTL 392

>PS00005|PD0C00005|PKC_PHOSPHO_SITE Protein kinase C phosphorylation site.

Query: 21 Query: 236 Query: 312 314 Query: 467 469 Query: 500 502

.

```
>PSGGGGG | PDOCGGGG | CK2_PHOSPHO_SITE Casein kinase II phosphorylation site.
Query: 3
               TSHE
Query: 258
               TLCD 261
>P300007 | PD0C00007 | TYR_PHOSPHO_SITE Tyrosine kinase phosphorylation site.
Query: 294
               RKGFEDWY
                               301
>PS66008 | PD0C00008 | MYRISTYL N-myristoylation site.
               GSQMAL 54
Ouerv: 49
Ouerv: 91
                GLFINT 96
Ouery: 98
                GCROTA 103
Query: 106
                GGLVNS 111
Query: 131
                GVAAGL 136
Query: 137
                GSGMAY 142
Query: 162
                GLSTTG 167
Query: 251
                GGLGNE 256
Query: 443
                GIIICA 448
>PS99929 | PD0C00029 | LEUCINE_ZIPPER Leucine zipper pattern.
Query: 387
                LANFTLVLSIFILPLMHTYAGL 408
>PS00342 | PDOC00299 | MICROBODIES_CTER Microbodies C-terminal targeting signal.
Query: 508
                AHV
                       510
```

Protein Family / Domain Matches, HMMer version 2

```
Searching for complete domains in PFAM humpfam - search a single seq against HOM database HOMER 2.1.1 (Dec 1998)
Copyright (C) 1992-1998 Washington University School of Medicine HOMER 2.1 freely distributed under the GNU General Public License (GPL).
HOMER is freely distributed under the GNU General Public License (GPL).
SOOM file: /prod/ddm/seqanal/PFAM/5fam5.5/Pfam
Sequence file: /prod/ddm/seqanal/PFAM/5fam5.5/Pfam
Sequence file: /prod/ddm/wspace/orfanal/oa-script.8673.seq
Parsed for domains:
Model Domain seq-f seq-t
                                             336 ..
477 ..
504 ..
Presenilin
 sugar_tr
Alignments of top-scoring domains:

Presentlin: domain 1 of 1, from 17 to 336: score -256.3, E = 4.8

*->eeelkkY:.....gakhvimifvPVsICHilVvsiks.vsfyeqnd

+ 1k +++ 4p + m++ 1 + ++++ e

25466 17 KDKKTLKPhpnidGGWAWMMVLSSFPVHILIMGSQWALGVLNVEWLE 63
                                  glkllYtpFhsdtptvgqklldSllNtlilIsVlvvmTfLlvvlYkyrFY
                             + Ph ++1 + +1 + 1+v f++++ ++
64 B------SRGLTAWVSSLSMGITLIVGPPIGLPINTCGCR 100.
             25466
                                   R...iIhgWLilSSlllLPlfsviyLqevfkaynDQDDNNQvp.....md
                          + iI.g +++Sl+ + + ++ +f ++ v+ +++m
101 QtaIIGG-LVNSLGWVLSAYAANVHYLFITFG-----VAaglgsgMA 141
             25466
                                   yptvllliwnPGvvGmmciHWkgpLrLqQfYLItisa....LmAlvfiky
                          y+ +++ ++ + + + L Q+ t + ++ Lm + +ky
142 YLPAVVMVGRY------PQKRRALAQGLSTTGTGfgtfLMTVL-LKY 181
             25466
                                   {\tt LPeWTaWfvLaaISvWDLVAVLtPkGPLrmLVBTAqeRnepiPPAliYSS}
                          L W mL A n ++ A1+
182 LCABYGWRN-------AMLIQGAVSLNLCVCGALMRPL 212
                          tyvvltvgsngeetnegtpsterrtskstiEYyTkReaQDDEFYQKIRQR
+ +++ e g p + ++kst
213 SPGKNPND-PGEKDVRGLPAHSTESVKST------------240
                                   rsaanPdsaptshst1VEAEPSpielkekssteeesddesdsseteGmPL + +++e+ d++ e++ L
```

http://lange.mpl.com/sequal/orfanal/25488.8679.htm

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vsgsSNLsssdssttvsqsdsllaphr.teeiareqpevqsnslpsNekr
               263 QA-----QECPDQAGHRKNMCALRILKTVS--WLTM---- 291
       25466
                    eeTAaDalhPdGeeErgvkLGLGDFiFYSvLvCkAsatasGdwnTtIACP
               V+ G+ D YS C AS t f
292 -----TNRMF 316
       25466
                    ValLiGLelTLlllavfkkALPALPisitfGlIFYFstdilvtPFmd<-*
               25466
MCT: domain 1 of 1, from 40 to 477: score 184.0, E = 2.5e-51
*->sflingfrdgfpksfgliffrelgeefqasnsetsvdsisSillav1
                       sf++++ '/+g + + g ++ +e++eef+ s++ t+w +ss+ ++ +
SPFVHILIMGSQMALG-VLNVEWLEEFHQSRGLTAW--VSSLSMGIT 83
       25466
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                 1++GP++ + n GcR++ i+Ggl+ s+g vl+ a n+ 1 +EfGv
84 LIVGPFIGLFINTCGCRQTAIIGGLVNSLGWVLSAYAANVHYLFITFGVA 133
       25466
                tGlGfafiylpaiviitsYpekkRsLAtgiavaGsgvgtfvlaplnpdqf
GlG ++ ylpaiv+++ Ypek+R+LA g++++G+g+gtf+ ++l+ ++
114 AGLGSGMAYLPAVVMVGRYFQKRRALAQGLSTTGTGFGTFLMTVLL--KY 181
       25466
                lienyGskWrgallffggmgyviAiwsvaivlncciagalfrplpsekvk

l + yG Wr+a+l+ g+ + ln+c++gal+rpl ++k

182 LCARYG--WRNAMLIQGA--------VSLNLCVCGALWRPLSPGK-- 216
                    qtklakaeepkealkskeneaseSidsirsaakaivspetpalslspelt
       +p 25466 217 ------ 218
              pkkdqlqkllktsrtrssngaklldfsvlkdargfllyassgslaslgtq
++++ + 1 +s++s g+q
219 -----NDPGEKDVRGLPAHSTESVKSTGQQ 243
       25466
                     lflpgsiflvnfakslgeslssvkskeaafllsilgdssdkegfggifar
       patllsflgfvanlketksnrpvliyllslcsivavvingilsrlasala
               25466
                     gsrkekkiksmidkielkstfwglflfslffgvgfgskkavvilalgfll
                + + 1f ++ f+ ++f ++a++ +

107 T-----ASLFTNRMFVAPIFW-----ALFAYSSFV 331
                fsilyaipvvglqkyssalGltetdAstlieaiavlniiGrplaGllaDk
+++ + +v l ++s+ ++ l+++ia++ i G +++G +aD
332 IPPIHLPEIVNLYNLSEQNDVF-----PLTSIIAIVHIFGKVILGVIADL 176
        25466
                     tknrklaiynlslilcGlfvafaPlatiflglafycvlfGsivfllayaF
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gf+ g y++l+++++dl+g + l na+g+++ + G+ +l Gpp+ag++
416 -GPSSG-YPSLMPVVTEDLVGIEHLANAYGIIICANGISALLGPPFACHI 463
        25466
                    kdltgsykvafyfa<-*
+d+t+ y+ sfy++
YDITQKYDFSFYIC 477
stvltglvvsifflGrliGslfaGklgdrfGRkksllialvlfviGalls

t++v's + li + f C +++ C + + +i+++ + C +ls
72 ----TAMVSSLSMGITLIVGPFIGLFINTCGRQTALIGGLVNSLGWVLS 117
        25466
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118 AYAANV-----HYLFITFGVAAGLGSGMAYLPAVVMVGRY-FQKRRA 158
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25466 302 SGYPGTASLFTN------RHFVAPIFWALFAYSSFVIPFIHLPEIVNL 343
                      .....vGvsdsrasllvtiivgvvNfvfTlvaliflvDrfGRRplli
                + +++++V +s ++ i++ +++ +++ +++ ++ ++ +1
364 ynlseqndvppLTS----IIAIVHIFGKVILGVIADLPCISV#---NVPL 386
        25466
                      lGaagmaicflilgasigvallllnkpkdplskaagivaivfillfiafP
               1 +++++ +i1 +1 ++ ++++++ F
387 LANFTLVLSIPIL-------PLMHTY-------AGLAVICALIG---P 417
                     algwCpipwvilsElPPtkvRskalalataanvlanfiigflfpyitgai
g+ ++ v++ +l + + a ++ +an +++ + + + + + i +
```

http://tange.mpi.com/seqansVortenat/25486,8670.htm

Analysis of 25466

```
25466 418 SSGYPSLMPVVTEDLVGIEHLANAYGIIICANGISALLGPPPAGWIYDIT 467
                                                        glalggyvflvfagllvlfilfvfffvPETKGrtLeeieelf<-*
y f ++ ++1 +++f+++ ++r +e+ ++
25466 468 Q----KYDFSPYI-CGLLYMIGILFLLIQPCIRIIEQSRRKY 504
//
Searching for complete domains in SMART
https:// Searching.com/SMART SMART 
      Scores for sequence family classification (score includes all domains):

Model Description Score E-value N

[no hits above thresholds]
      Parsed for domains:
Nodel Domain seq-f seq-t hmm-f hmm-t

[no hits above thresholds]
```

score E-value

ProDom Matches

Alignments of top-scoring domains: [no hits above thresholds]

ProdomId	Start	End	Description	Score
View Prodom 38068 (BROX 학생 (단) (Showing match 항목가 함복하는)	345	461	p99.2 (2) 006142(1) 030820(1) // PROTEIN ADENYLYL CYCLASE	73
View Prodom 102799 (Blover 1971) (Showing match 1972) (God)	393	467	p99.2 (1) YU45_MYCTU // HYPOTHETICAL 41.9 KD PROTEIN CY339.45C TRANSMEMBRANE :	80
View Prodom 84838 [RBöxer 2004] Schöwing match (1992-1993) [4]	440	494	p99.2 (1) Q23113_CAEEL // SIMILAR TO SYNAPTIC VESICLE AMINE TRANSPORTER	74
ProdomId	Start	End	Description	Score

>84838 p99.2 (1) Q23113_CAEEL // SIMILAR TO SYNAPTIC VESICLE AMINE TRANSPORTER Length = 85 Score = 74 (31.1 bits), Expect = 0.053, P = 0.052 Identities = 18/57 (31%), Positives = 31/57 (54%)

Query: 440 NAYGIIICANGISALLGPPPAGMIYDITOKYDPSFYICGLLYHI--GILFLLIOPCI 494
+ # GI + L4GPP+ G * Y + K + F + LL + I P+++QF II
14 SAMGIALGGLALGVLVGPYGGLIKYMSG-ELPPYALLALLALPDGSIGPYALQFKI 69

View Prodom 102799

>102799 p99.2 (1) YU45_MYCTU // HYPOTHETICAL 41.9 KD PROTEIN CY339.45C TRANSMEMBRANE Length = 155

Score = 80 (33.2 bits), Expect = 0.55, P = 0.43 Identities = 25/78 (32%), Positives = 37/78 (47%)

Query: 193 VLSIFILPLMHT-YAGLAVICALI-GFSSGYF-SLMPVVTEDLVGIBHLANAYGIIICAN 449
+LSI++ T + VI L+ C + G S + V DLV A+AYG+
Sbjct: 48 ILSILVVLFAPTDNVTMVVIGTLVMGAAVGIQESTLRGVVADLVASPRRASAYGVFAAGL 107

Query: 450 GISALLGPPFAGWIYDIT 467 Query: 450 GISALLGPFFAGRIDE: C + G GW+YDI+
Sbjct: 108 GAATAGGGALIGWLYDIS 125

View Prodom 38068

>38068 p99.2 (2) 006142(1) 030820(1) // PROTEIN ADENYLYL CYCLASE Length = 218

Score = 73 (30.8 bits), Expect = 8.1, P = 1.0

http://tango.mpi.com/seqanaVortanal/25466.6670.html

Identities = 41/135 (30%), Positives = 62/135 (45%)

Query:	345 NLSEQNDVFPLTSIIAIVHIFGKVILGVIADLPCISV#NVFLLANPTLVLSIF 397
	15 SVARRORVLTITAWLAVVVTGSFALMQLATGAGGWYIALINVPTAVTFAIVPLLHRFGGL 74
Sbjct:	15 SVARKORVETTTAWEAVVVIGSFALMODATOAGGMTTABLITTT INVITATIVE BERNINGS
Query:	398 ILPLMHTYAGLAVICAL-IGPSSGYFSLMPVVTEDLVGIEHLANAYGIIICAN 44
	+ PL T+ G A + IG+ G +F + LVGIEH A A G+ A
Sbjct:	75 VAPLTPIGTAYVAIFAIGWDVGTDAGAQFFFLVAAALVVLLVGIEHTALAVGLAAVAA 113
Query:	450 G-ISALLGPPPAG 461
	G + AL L PP G

Query: 450 G-ISAL--LGPPPAG 461 G + AL L PP G Sbjet: 133 GLVIALEFLVPPDTG 147

Exhibit D to Accompany Declaration under 37 CFR §1.131 for US Appln. No. 10/074,547

Fbh25466pat 015374	MYTSHEDIGYDFEDGPKDKKTLKPHPNIDGGWAWMMVLSSFFVHILIMGSQMALGVLNVEMLKRE-GKVQPYTKTLDGGWGWMIVIHFFLVNVFVMGMTKTFAIFFVV . * * : * .:***.**: *:*:::** ::::: *
Fbh25466pat 015374	WLEEFHQSRGLTAWVSSLSMGITLIVGPFIGLFINTCGCRQTAIIGGLVNSLGWVLSAYA FQEEFEGTSEQIGWIGSIMSSLRFCAGPLVAIICDILGEKTTSILGAFVVTGGYLISSWA : ***.: .*:.*: .: : .**:.:: : * : *:*:*:* : *::.*:
Fbh25466pat 015374	ANVHYLFITFGVXAGLGSGMAYLPAVXMVGRYFQKRRALAQGLSTTGTGFGTFLMTVLLK TSIPFLCVTMGLLPGLGSAFLYQVAAVVTTKYFKKRLALSTAIARSGMGL-TFLLAPFTK :.: :* :*:*: ****: : ****: : ** * * : : * : *
Fbh25466pat 015374	YLCAEYGWRNAMLIQGAVSLNLCVCGALMRPLSPGKNPNDPGEKDVRG-LPAHSTESVKS FLIDLYDWTGALILFGAIALNLVPSSMLLRPI-HIKSENNSGIKDKGSSLSAHGPEAHAT :* *.*.*:: **::*** *:**: *. *:.* ** . *.***: :
Fbh25466pat 015374	TGQQGRTEEKDGGLGNE-ETLCDLQAQECPDQAGHRKNMCALRILKTVSWLTMR ETHCHETEESTIKDSTTQKAGLPSKNLTVSQNQSEEFYNGPNRNRLLLKSDEESDK : .*** **. * :.* * *.: * :: :: :: ::
Fbh25466pat 015374	VRKGFEDWYSGYFGTASLFTNRMFVAFIFWA-LFAYSSFVIPFIHLPEIVNLYNLSEQND VISWSCKQLFDISLFRNPFFYIFT-WSFLLSQLAYFIPTFHLVARAKTLGID-IMD * . * . : *** * : * .: .:. *
Fbh25466pat 015374	VFPLTSIIAIVHIFGKVILGVIADLPCISVWNVFLLANFTLVLSIFILPLMHTYAGLA ASYLVSVAGILETVSQIISGWVADQNWIKKYHYHKSYLILCGITNLLAPLATTFPLLMTY . *.*: .*: . : : : * * : . * * * * *
Fbh25466pat 015374	VICALIGFSSGYFSLMPVVTEDLVGIEHLANAYGIIICANGISALLGPPFAGWIYDITQK TICFAI-FAGGYLALILPVLVDLCRNSTVNRFLGLASFFAGMAVLSGPPIAGWLYDYTQT .** * *:.**::*: * ** . : . *: *::.* ***:** **.
Fbh25466pat 015374	YDFSFYICGLLYMIGILFLLIQPCIRIIEQSRRKYMDGAHV YNGSFYFSGICYLLSSVSFFFVPLAERWKNSLT *: ***: *: : : : : : : : : : : : : : :